SEQUENCE LISTING

- <110> Rudiger Ridder, et.al.
- Method for solution based diagnosis <120>
- <130> 05033.0003.00us00
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- 10/633,484 2003-07-31 <141>
- <150> EP 02017313.4
- <151> 2002-08-01
- <160> 14
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- Cys Gly Arg Gln Tyr Thr Leu Lys Lys Thr Thr Thr Tyr Thr Gln Gly 50 60
- Val Pro Pro Ser Gln Gly Asp Leu Glu Tyr Gln Met Ser Thr Thr Ala 65 70 75 80
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400

Ser Val Asp Asp Val Asn Val Leu Thr Cys Ala Thr Gly Thr Leu Ser $405 \hspace{1.5cm} 410 \hspace{1.5cm} 415$

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Asn Leu Thr Cys Asn Asn Ser Lys Asn Lys Thr Leu Val Thr Gln Asn 420 425 430

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Asp Asp Ile Thr Glu Pro Ala Val Cys Ala Leu Arg His Leu Thr Ser 450 455 460

Arg His Pro Glu Ala Glu Met Ala Gln Asn Ser Val Arg Leu Asn Tyr 465 470 475 480

Gly Ile Pro Ala Ile Val Lys Leu Leu Asn Gln Pro Asn Gln Trp Pro 485 490 495

Leu Val Lys Ala Thr Ile Gly Leu Ile Arg Asn Leu Ala Leu Cys Pro 500 505 510

Ala Asn His Ala Pro Leu Gln Glu Ala Ala Val Ile Pro Arg Leu Val

Gln Leu Leu Val Lys Ala His Gln Asp Ala Gln Arg His Val Ala Ala 530 535 540

Gly Thr Gln Gln Pro Tyr Thr Asp Gly Val Arg Met Glu Glu Ile Val 545 550 555 560

Glu Gly Cys Thr Gly Ala Leu His Ile Leu Ala Arg Asp Pro Met Asn 565 570 575

Arg Met Glu Ile Phe Arg Leu Asn Thr Ile Pro Leu Phe Val Gln Leu 580 585 590

Leu Tyr Ser Ser Val Glu Asn Ile Gln Arg Val Ala Ala Gly Val Leu 595 600 605

Cys Glu Leu Ala Gln Asp Lys Glu Ala Ala Asp Ala Ile Asp Ala Glu 610 620

Gly Ala Ser Ala Pro Leu Met Glu Leu Leu His Ser Arg Asn Glu Gly 625 630 635 640

Thr Ala Thr Tyr Ala Ala Ala Val Leu Phe Arg Ile Ser Glu Asp Lys Page 3

645 650 655

Asn Pro Asp Tyr Arg Lys Arg Val Ser Val Glu Leu Thr Asn Ser Leu 660 665 670

Phe Lys His Asp Pro Ala Ala Trp Glu Ala Ala Gln Ser Met Ile Pro 675 680 685

Ile Asn Glu Pro Tyr Gly Asp Asp Leu Asp Ala Thr Tyr Arg Pro Met 690 695 700

Tyr Ser Ser Asp Val Pro Leu Asp Pro Leu Glu Met His Met Asp Met 705 715 720

Asp Gly Asp Tyr Pro Ile Asp Thr Tyr Ser Asp Gly Leu Arg Pro Pro 725 730 735

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Thr Ser Val Gly Ala Gln Asn Thr Val Ile Cys Ser Lys Leu Ala Ala 50 60

Lys Cys Leu Val Met Lys Ala Glu Met Asn Gly Ser Lys Leu Gly Arg 65 70 75 80

Arg Ala Lys Pro Glu Gly Ala Leu Gln Asn Asn Asp Gly Leu Tyr Asp 85 90 95 Pro Asp Cys Asp Glu Ser Gly Leu Phe Lys Ala Lys Gln Cys Asn Gly 100 105 110Thr Ser Thr Cys Trp Cys Val Asn Thr Ala Gly Val Arg Arg Thr Asp 115 120 125 Lys Asp Thr Glu Ile Thr Cys Ser Glu Arg Val Arg Thr Tyr Trp Ile 130 135 140 Ile Ile Glu Leu Lys His Lys Ala Arg Glu Lys Pro Tyr Asp Ser Lys 145 150 155 160 Ser Leu Arg Thr Ala Leu Gln Lys Glu Ile Thr Thr Arg Tyr Gln Leu 165 170 175 Asp Pro Lys Phe Ile Thr Ser Ile Leu Tyr Glu Asn Asn Val Ile Thr 180 185 190 Ile Asp Leu Val Gln Asn Ser Ser Gln Lys Thr Gln Asn Asp Val Asp 195 200 205 Ile Ala Asp Val Ala Tyr Tyr Phe Glu Lys Asp Val Lys Gly Glu Ser 210 215 220 Leu Phe His Ser Lys Lys Met Asp Leu Thr Val Asn Gly Glu Gln Leu 225 230 235 240 Asp Leu Asp Pro Gly Gln Thr Leu Ile Tyr Tyr Val Asp Glu Lys Ala 245 250 255 Pro Glu Phe Ser Met Gln Gly Leu Lys Ala Gly Val Ile Ala Val Ile 260 265 270Val Val Val Met Ala Val Val Ala Gly Ile Val Val Leu Val Ile Ser Arg Lys Lys Arg Met Ala Lys Tyr Glu Lys Ala Glu Ile Lys Glu 290 295 300 Met Gly Glu Met His Arg Glu Leu Asn Ala 305 310

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Ile Ala Thr Tyr Thr Leu Phe Ser His Ala Val Ser Ser Asn Gly Asn 225 230 235 240 Ala Val Glu Asp Pro Met Glu Ile Leu Ile Thr Val Thr Asp Gln Asn 245 250 255 Asp Asn Lys Pro Glu Phe Thr Gln Glu Val Phe Lys Gly Ser Val Met 260 265 270 Glu Gly Ala Leu Pro Gly Thr Ser Val Met Glu Val Thr Ala Thr Asp 275 280 285 Ala Asp Asp Asp Val Asn Thr Tyr Asn Ala Ile Ala Tyr Thr Ile 290 295 300 Leu Ser Gln Asp Pro Glu Leu Pro Asp Lys Asn Met Phe Thr Ile Asn 305 310 315 320 Arg Asn Thr Gly Val Ile Ser Val Val Thr Thr Gly Leu Asp Arg Glu 325 330 335 Ser Phe Pro Thr Tyr Thr Leu Val Val Gln Ala Ala Asp Leu Gln Gly Glu Gly Leu Ser Thr Thr Ala Thr Ala Val Ile Thr Val Thr Asp Thr Asn Asp Asn Pro Pro Ile Phe Asn Pro Thr Tyr Lys Gly Gln Val 370 375 380 Pro Glu Asn Glu Ala Asn Val Val Ile Thr Thr Leu Lys Val Thr Asp 395 Ala Asp Ala Pro Asn Thr Pro Ala Trp Glu Ala Val Tyr Thr Ile Leu Asn Asp Asp Gly Gly Gln Phe Val Val Thr Thr Asn Pro Val Asn Asn 420 425 430 Asp Gly Ile Leu Lys Thr Ala Lys Gly Leu Asp Phe Glu Ala Lys Gln 445 Gln Tyr Ile Leu His Val Ala Val Thr Asn Val Val Pro Phe Glu Val 450 455 460 Ser Leu Thr Thr Ser Thr Ala Thr Val Thr Val Asp Val Leu Asp Val 465 470 475 480 480 Asn Glu Ala Pro Ile Phe Val Pro Pro Glu Lys Arg Val Glu Val Ser Glu Asp Phe Gly Val Gly Gln Glu Ile Thr Ser Tyr Thr Ala Gln Glu 500 510 Pro Asp Thr Phe Met Glu Gln Lys Ile Thr Tyr Arg Ile Trp Arg Asp 515 520 525 Thr Ala Asn Trp Leu Glu Ile Asn Pro Asp Thr Gly Ala Ile Ser Thr 530 535 540 Arg Ala Glu Leu Asp Arg Glu Asp Phe Glu His Val Lys Asn Ser Thr 545 550 555 560 Tyr Thr Ala Leu Ile Ile Ala Thr Asp Asn Gly Ser Pro Val Ala Thr 565 570 575 Gly Thr Gly Thr Leu Leu Leu Ile Leu Ser Asp Val Asn Asp Asn Ala 580 585 Pro Ile Pro Glu Pro Arg Thr Ile Phe Phe Cys Glu Arg Asn Pro Lys 595 600 605 Pro Gln Val Ile Asn Ile Ile Asp Ala Asp Leu Pro Pro Asn Thr Ser 610 620 Pro Phe Thr Ala Glu Leu Thr His Gly Ala Ser Ala Asn Trp Thr Ile Gln Tyr Asn Asp Pro Thr Gln Glu Ser Ile Ile Leu Lys Pro Lys Met 645 650 655 Ala Leu Glu Val Gly Asp Tyr Lys Ile Asn Leu Lys Leu Met Asp Asn 660 670 Gln Asn Lys Asp Gln Val Thr Thr Leu Glu Val Ser Val Cys Asp Cys 685 Glu Gly Ala Ala Gly Val Cys Arg Lys Ala Gln Pro Val Glu Ala Gly 690 695 700 Leu Gln Ile Pro Ala Ile Leu Gly Ile Leu Gly Gly Ile Leu Ala Leu 705 710 715 720 Leu Ile Leu Ile Leu Leu Leu Leu Phe Leu Arg Arg Ala Val 725 730 735 Val Lys Glu Pro Leu Leu Pro Pro Glu Asp Asp Thr Arg Asp Asn Val 740 745 750

Tyr Tyr Asp Glu Glu Gly Gly Glu Glu Asp Gln Asp Phe Asp 755 760 765 Leu Ser Gln Leu His Arg Gly Leu Asp Ala Arg Pro Glu Val Thr Arg 770 780 Asn Asp Val Ala Pro Thr Leu Met Ser Val Pro Arg Tyr Leu Pro Arg 785 790 795 800 Pro Ala Asn Pro Asp Glu Ile Gly Asn Phe Ile Asp Glu Asn Leu Lys 805 810 815 Ala Ala Asp Thr Asp Pro Thr Ala Pro Pro Tyr Asp Ser Leu Leu Val 820 825 830 Phe Asp Tyr Glu Gly Ser Gly Ser Glu Ala Ala Ser Leu Ser Ser Leu 835 840 845 Asn Ser Ser Glu Ser Asp Lys Asp Gln Asp Tyr Asp Tyr Leu Asn Glu 850 860 Trp Gly Asn Arg Phe Lys Lys Leu Ala Asp Met Tyr Gly Gly Glu 865 870 880 Asp Asp <210> <211> 906 **PRT** Homo sapiens <220> misc_feature
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Val Glu Gln Ala Thr Glu Asn Phe Leu Glu Lys Gly Asp Lys Ile Ala 65 70 75 80 Lys Glu Ser Gln Phe Leu Lys Glu Glu Leu Val Ala Ala Val Glu Asp 85 90 95 Val Arg Lys Gln Gly Asp Leu Met Lys Ala Ala Ala Gly Glu Phe Ala 100 105 110 Asp Asp Pro Cys Ser Ser Val Lys Arg Gly Asn Met Val Arg Ala Ala 115 120 125 Arg Ala Leu Leu Ser Ala Val Thr Arg Leu Leu Ile Leu Ala Asp Met Ala Asp Val Tyr Lys Leu Leu Val Gln Leu Lys Val Val Glu Asp Gly 145 150 155 160 Ile Leu Lys Leu Arg Asn Ala Gly Asn Glu Gln Asp Leu Gly Ile Gln 165 170 175 Tyr Lys Ala Leu Lys Pro Glu Val Asp Lys Leu Asn Ile Met Ala Ala 180 185 190 Lys Arg Gln Glu Leu Lys Asp Val Gly His Arg Asp Gln Met Ala 195 200 205 Ala Ala Arg Gly Ile Leu Gln Lys Asn Val Pro Ile Leu Tyr Thr Ala 210 215 220 Ser Gln Ala Cys Leu Gln His Pro Asp Val Ala Ala Tyr Lys Ala Asn 225 230 235 240 Arg Asp Leu Ile Tyr Lys Gln Leu Gln Gln Ala Val Thr Gly Ile Ser 245 250 255 Asn Ala Ala Gln Ala Thr Ala Ser Asp Asp Ala Ser Gln His Gln Gly 260 265 270 Gly Gly Gly Glu Leu Ala Tyr Ala Leu Asn Asn Phe Asp Lys Gln 280 285 Ile Ile Val Asp Pro Leu Ser Phe Ser Glu Glu Arg Phe Arg Pro Ser 290 295 300 Page 10

Asp Ser Ser Cys Thr Arg Asp Asp Arg Glu Arg Ile Val Ala Glu 325 330 335 Cys Asn Ala Val Arg Gln Ala Leu Gln Asp Leu Leu Ser Glu Tyr Met Gly Asn Ala Gly Arg Lys Glu Arg Ser Asp Ala Leu Asn Ser Ala Ile 355 360 365 Asp Lys Met Thr Lys Lys Thr Arg Asp Leu Arg Arg Gln Leu Arg Lys 370 380 Ala Val Met Asp His Val Ser Asp Ser Phe Leu Glu Thr Asn Val Pro Leu Leu Val Leu Ile Glu Ala Ala Lys Asn Gly Asn Glu Lys Glu Val Lys Glu Tyr Ala Gln Val Phe Arg Glu His Ala Asn Lys Leu Ile Glu Val Ala Asn Leu Ala Cys Ser Ile Ser Asn Asn Glu Glu Gly Val Lys Leu Val Arg Met Ser Ala Ser Gln Leu Glu Ala Leu Cys Pro Gln Val Ile Asn Ala Ala Leu Ala Leu Ala Ala Lys Pro Gln Ser Lys Leu Ala Gln Glu Asn Met Asp Leu Phe Lys Glu Gln Trp Glu Lys Gln Val Arg Val Leu Thr Asp Ala Val Asp Asp Ile Thr Ser Ile Asp Asp Phe Leu 500 510 Ala Val Ser Glu Asn His Ile Leu Glu Asp Val Asn Lys Cys Val Ile 515 520 525 Ala Leu Gln Glu Lys Asp Val Asp Gly Leu Asp Arg Thr Ala Gly Ala 530 540 Ile Arg Gly Arg Ala Ala Arg Val Ile His Val Val Thr Ser Glu Met Asp Asn Tyr Glu Pro Gly Val Tyr Thr Glu Lys Val Leu Glu Ala Thr Page 11

Lys Leu Leu Ser Asn Thr Val Met Pro Arg Phe Thr Glu Gln Val Glu Ala Ala Val Glu Ala Leu Ser Ser Asp Pro Ala Gln Pro Met Asp Glu Asn Glu Phe Ile Asp Ala Ser Arg Leu Val Tyr Asp Gly Ile Arg Asp 610 615 620 Ile Arg Lys Ala Val Leu Met Ile Arg Thr Pro Glu Glu Leu Asp Asp Ser Asp Phe Glu Thr Glu Asp Phe Asp Val Arg Ser Arg Thr Ser Val 645 650 655 Gln Thr Glu Asp Asp Gln Leu Ile Ala Gly Gln Ser Ala Arg Ala Ile 660 665 670 Met Ala Gln Leu Pro Gln Glu Gln Lys Ala Lys Ile Ala Glu Gln Val 675 680 685 Ala Ser Phe Gln Glu Glu Lys Ser Lys Leu Asp Ala Glu Val Ser Lys 690 695 700 Trp Asp Asp Ser Gly Asn Asp Ile Ile Val Leu Ala Lys Gln Met Cys 705 710 715 720 Met Ile Met Met Glu Met Thr Asp Phe Thr Arg Gly Lys Gly Pro Leu 725 730 735 Lys Asn Thr Ser Asp Val Ile Ser Ala Ala Lys Lys Ile Ala Glu Ala 740 745 750 Gly Ser Arg Met Asp Lys Leu Gly Arg Thr Ile Ala Asp His Cys Pro 755 760 765 Asp Ser Ala Cys Lys Gln Asp Leu Leu Ala Tyr Leu Gln Arg Ile Ala 770 780 Leu Tyr Cys His Gln Leu Asn Ile Cys Ser Lys Val Lys Ala Glu Val 785 790 795 800 Gln Asn Leu Gly Gly Glu Leu Val Val Ser Gly Val Asp Ser Ala Met 805 810 815 Ser Leu Ile Gln Ala Ala Lys Asn Leu Met Asn Ala Val Val Gln Thr 820 825 830 Page 12

Val Lys Ala Ser Tyr Val Ala Ser Thr Lys Tyr Gln Lys Ser Gln Gly 845

Met Ala Ser Leu Asn Leu Pro Ala Val Ser Trp Lys Met Lys Ala Pro 860

Glu Lys Lys Pro Leu Val Lys Arg Glu Lys Gln Asp Glu Thr Gln Thr 865 870 875

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Homo sapiens

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Lys Lys Lys Gly Arg Ser Lys Lys Ala His Val Leu Ala Ala Ser Val 50 60

Glu Gln Ala Thr Gln Asn Phe Leu Glu Lys Gly Glu Gln Ile Ala Lys 65 70 75 80

Glu Ser Gln Asp Leu Lys Glu Glu Leu Val Ala Ala Val Glu Asp Val 85 90 95

Arg Lys Gln Gly Glu Thr Met Arg Ile Ala Ser Ser Glu Phe Ala Asp $100 \hspace{1.5cm} 105 \hspace{1.5cm} 110$

Asp Pro Cys Ser Ser Val Lys Arg Gly Thr Met Val Arg Ala Ala Arg Page 13

115 120 125

Ala Leu Leu Ser Ala Val Thr Arg Leu Leu Ile Leu Ala Asp Met Ala 130 135 140 Asp Val Met Arg Leu Leu Ser His Leu Lys Ile Val Glu Glu Ala Leu 145 150 155 160 Glu Ala Val Lys Asn Ala Thr Asn Glu Gln Asp Leu Ala Asn Arg Phe 165 170 175 Lys Glu Phe Gly Lys Glu Met Val Lys Leu Asn Tyr Val Ala Ala Arg 180 185 190 Arg Gln Gln Glu Leu Lys Asp Pro His Cys Arg Asp Glu Met Ala Ala 195 200 205 Ala Arg Gly Ala Leu Lys Lys Asn Ala Thr Met Leu Tyr Thr Ala Ser 210 215 220 Gln Ala Phe Leu Arg His Pro Asp Val Ala Ala Thr Arg Ala Asn Arg 225 230 235 240 Asp Tyr Val Phe Lys Gln Val Gln Glu Ala Ile Ala Gly Ile Ser Asn 245 250 255 Ala Ala Gln Ala Thr Ser Pro Thr Asp Glu Ala Lys Gly His Thr Gly 260 265 270Ile Gly Glu Leu Ala Ala Ala Leu Asn Glu Phe Asp Asn Lys Ile Ile 275 280 285 Leu Asp Pro Met Thr Phe Ser Glu Ala Arg Phe Arg Pro Ser Leu Glu 290 295 300 Glu Arg Leu Glu Ser Ile Ile Ser Gly Ala Ala Leu Met Ala Asp Ser 305 315 320 Ser Cys Thr Arg Asp Asp Arg Glu Arg Ile Val Ala Glu Cys Asn 325 330 335 Ala Val Arg Gln Ala Leu Gln Asp Leu Leu Ser Glu Tyr Met Asn Asn 340 350 Thr Gly Arg Lys Glu Lys Gly Asp Pro Leu Asn Ile Ala Ile Asp Lys 355 360 365 Met Thr Lys Lys Thr Arg Asp Leu Arg Arg Gln Leu Arg Lys Ala Val

Met Asp His Ile Ser Asp Ser Phe Leu Glu Thr Asn Val Pro Leu Leu 385 390 400 Val Leu Ile Glu Ala Ala Lys Ser Gly Asn Glu Lys Glu Val Lys Glu 405 410 415 Tyr Ala Gln Val Phe Arg Glu His Ala Asn Lys Leu Val Glu Val Ala 420 425 430 Asn Leu Ala Cys Ser Ile Ser Asn Asn Glu Glu Gly Val Lys Leu Val 435 440 445 Arg Met Ala Ala Thr Gln Ile Asp Ser Leu Cys Pro Gln Val Ile Asn 450 460 Ala Ala Leu Thr Leu Ala Ala Arg Pro Gln Ser Lys Val Ala Gln Asp 465 470 475 480 Asn Met Asp Val Phe Lys Asp Gln Trp Glu Lys Gln Val Arg Val Leu Thr Glu Ala Val Asp Asp Ile Thr Ser Val Asp Asp Phe Leu Ser Val 500 505 510Ser Glu Asn His Ile Leu Glu Asp Val Asn Lys Cys Val Ile Ala Leu 515 520 525 Gln Glu Gly Asp Val Asp Thr Leu Asp Arg Thr Ala Gly Ala Ile Arg 530 535 540 Gly Arg Ala Ala Arg Val Ile His Ile Ile Asn Ala Glu Met Glu Asn 545 550 560 Tyr Glu Ala Gly Val Tyr Thr Glu Lys Val Leu Glu Ala Thr Lys Leu
565 570 575 Leu Ser Glu Thr Val Met Pro Arg Phe Ala Glu Gln Val Glu Val Ala Ile Glu Ala Leu Ser Ala Asn Val Pro Gln Pro Phe Glu Glu Asn Glu Phe Ile Asp Ala Ser Arg Leu Val Tyr Asp Gly Val Arg Asp Ile Arg 610 615 620 Lys Ala Val Leu Met Ile Arg Thr Pro Glu Glu Leu Glu Asp Asp Ser

Asp Phe Glu Gln Glu Asp Tyr Asp Val Arg Ser Arg Thr Ser Val Gln 645 650 655 Thr Glu Asp Asp Gln Leu Ile Ala Gly Gln Ser Ala Arg Ala Ile Met
660 665 670 Ala Gln Leu Pro Gln Glu Glu Lys Ala Lys Ile Ala Glu Gln Val Glu 675 680 685 Ile Phe His Gln Glu Lys Ser Lys Leu Asp Ala Glu Val Ala Lys Trp 690 695 700 Asp Asp Ser Gly Asn Asp Ile Ile Val Leu Ala Lys Gln Met Cys Met 705 710 715 720 Ile Met Met Glu Met Thr Asp Phe Thr Arg Gly Lys Gly Pro Leu Lys 725 730 735 Asn Thr Ser Asp Val Ile Asn Ala Ala Lys Lys Ile Ala Glu Ala Gly 740 745 750 Ser Arg Met Asp Lys Leu Ala Arg Ala Val Ala Asp Gln Cys Pro Asp 765 760 765 Ser Ala Cys Lys Gln Asp Leu Leu Ala Tyr Leu Gln Arg Ile Ala Leu 770 775 780 Tyr Cys His Gln Leu Asn Ile Cys Ser Lys Val Lys Ala Glu Val Gln 785 790 795 800 Asn Leu Gly Gly Glu Leu Ile Val Ser Gly Thr Gly Val Gln Ser Thr 805 810 815 Phe Thr Thr Phe Tyr Glu Val Asp Cys Asp Val Ile Asp Gly Gly Arg 820 825 Ala Ser Gln Leu Ser Thr His Leu Pro Thr Cys Ala Glu Gly Ala Pro 835 840 845 Ile Gly Ser Gly Ser Ser Asp Ser Ser Met Leu Asp Ser Ala Thr Ser 850 855 860 Leu Ile Gln Ala Ala Lys Asn Leu Met Asn Ala Val Val Leu Thr Val 865 870 875 880 880 Lys Ala Ser Tyr Val Ala Ser Thr Lys Tyr Gln Lys Val Tyr Gly Thr

Ala Ala Val Asn Ser Pro Val Val Ser Trp Lys Met Lys Ala Pro Glu 905

Lys Lys Pro Leu Val Lys Arg Glu Lys Pro Glu Glu Phe Gln Thr Arg 915 920 925

Val Arg Arg Gly Ser Gln Lys Lys His Ile Ser Pro Val Gln Ala Leu 930 935 940

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beta-Catenin, Swissprot Accession P35222

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Ser Gly Ile His Ser Gly Ala Thr Thr Ala Pro Ser Leu Ser Gly 35 40 45

Lys Gly Asn Pro Glu Glu Glu Asp Val Asp Thr Ser Gln Val Leu Tyr 50 55 60

Glu Trp Glu Gln Gly Phe Ser Gln Ser Phe Thr Gln Glu Gln Val Ala 65 70 75 80

Asp Ile Asp Gly Gln Tyr Ala Met Thr Arg Ala Gln Arg Val Arg Ala 85 90 95

Ala Met Phe Pro Glu Thr Leu Asp Glu Gly Met Gln Ile Pro Ser Thr

Gln Phe Asp Ala Ala His Pro Thr Asn Val Gln Arg Leu Ala Glu Pro 115 120 125

Ser Gln Met Leu Lys His Ala Val Val Asn Leu Ile Asn Tyr Gln Asp 130 135 140 Asp Ala Glu Leu Ala Thr Arg Ala Ile Pro Glu Leu Thr Lys Leu Leu 145 150 160 Asn Asp Glu Asp Gln Val Val Val Asn Lys Ala Ala Val Met Val His 165 170 175 Gln Leu Ser Lys Lys Glu Ala Ser Arg His Ala Ile Met Arg Ser Pro 180 185 190 Gln Met Val Ser Ala Ile Val Arg Thr Met Gln Asn Thr Asn Asp Val 195 200 205 Glu Thr Ala Arg Cys Thr Ala Gly Thr Leu His Asn Leu Ser His His 210 215 220 Arg Glu Gly Leu Leu Ala Ile Phe Lys Ser Gly Gly Ile Pro Ala Leu 225 230 235 240 Val Lys Met Leu Gly Ser Pro Val Asp Ser Val Leu Phe Tyr Ala Ile 245 250 255 Thr Thr Leu His Asn Leu Leu Leu His Gln Glu Gly Ala Lys Met Ala 260 265 270 Val Arg Leu Ala Gly Gly Leu Gln Lys Met Val Ala Leu Leu Asn Lys 275 280 285 Thr Asn Val Lys Phe Leu Ala Ile Thr Thr Asp Cys Leu Gln Ile Leu 290 295 300 Ala Tyr Gly Asn Gln Glu Ser Lys Leu Ile Ile Leu Ala Ser Gly Gly 305 310 315 320 Pro Gln Ala Leu Val Asn Ile Met Arg Thr Tyr Thr Tyr Glu Lys Leu 325 330 335 Leu Trp Thr Thr Ser Arg Val Leu Lys Val Leu Ser Val Cys Ser Ser 340 345 350 Asn Lys Pro Ala Ile Val Glu Ala Gly Gly Met Gln Ala Leu Gly Leu 355 360 365 His Leu Thr Asp Pro Ser Gln Arg Leu Val Gln Asn Cys Leu Trp Thr 370 375 380 Leu Arg Asn Leu Ser Asp Ala Ala Thr Lys Gln Glu Gly Met Glu Gly 385 390 395 Leu Leu Gly Thr Leu Val Gln Leu Leu Gly Ser Asp Asp Ile Asn Val 405 410 415 Val Thr Cys Ala Ala Gly Ile Leu Ser Asn Leu Thr Cys Asn Asn Tyr 420 425 430 Lys Asn Lys Met Met Val Cys Gln Val Gly Gly Ile Glu Ala Leu Val 435 440 445 Arg Thr Val Leu Arg Ala Gly Asp Arg Glu Asp Ile Thr Glu Pro Ala 450 460 Ile Cys Ala Leu Arg His Leu Thr Ser Arg His Gln Glu Ala Glu Met 465 470 475 480 Ala Gln Asn Ala Val Arg Leu His Tyr Gly Leu Pro Val Val Lys 485 490 495 Leu Leu His Pro Pro Ser His Trp Pro Leu Ile Lys Ala Thr Val Gly 500 510 Leu Ile Arg Asn Leu Ala Leu Cys Pro Ala Asn His Ala Pro Leu Arg 515 520 525 Glu Gln Gly Ala Ile Pro Arg Leu Val Gln Leu Leu Val Arg Ala His 530 535 Gln Asp Thr Gln Arg Arg Thr Ser Met Gly Gly Thr Gln Gln Gln Phe 545 550 555 Val Glu Gly Val Arg Met Glu Glu Ile Val Glu Gly Cys Thr Gly Ala 565 570 575 Leu His Ile Leu Ala Arg Asp Val His Asn Arg Ile Val Ile Arg Gly Leu Asn Thr Ile Pro Leu Phe Val Gln Leu Leu Tyr Ser Pro Ile Glu Asn Ile Gln Arg Val Ala Ala Gly Val Leu Cys Glu Leu Ala Gln Asp 610 615 620 Lys Glu Ala Ala Glu Ala Ile Glu Ala Glu Gly Ala Thr Ala Pro Leu 625 635 640 Thr Glu Leu Leu His Ser Arg Asn Glu Gly Val Ala Thr Tyr Ala Ala 645 650 655 Ala Val Leu Phe Arg Met Ser Glu Asp Lys Pro Gln Asp Tyr Lys Lys 660 665 670 Arg Leu Ser Val Glu Leu Thr Ser Ser Leu Phe Arg Thr Glu Pro Met 675 680 685 Ala Trp Asn Glu Thr Ala Asp Leu Gly Leu Asp Ile Gly Ala Gln Gly 690 695 700 Glu Pro Leu Gly Tyr Arg Gln Asp Asp Pro Ser Tyr Arg Ser Phe His 705 715 720 Ser Gly Gly Tyr Gly Gln Asp Ala Leu Gly Met Asp Pro Met Met Glu
725 730 735 His Glu Met Gly Gly His His Pro Gly Ala Asp Tyr Pro Val Asp Gly 740 745 750 Leu Pro Asp Leu Gly His Ala Gln Asp Leu Met Asp Gly Leu Pro Pro 765 760 765 Gly Asp Ser Asn Gln Leu Ala Trp Phe Asp Thr Asp Leu 770 780 <210> 585 <211> <212> PRT <213> Homo sapiens <220> <221> misc_feature (1). (585) Involucrin, Swissprot Accession P07476 <400> 7 Met Ser Gln Gln His Thr Leu Pro Val Thr Leu Ser Pro Ala Leu Ser 10 15Gln Glu Leu Leu Lys Thr Val Pro Pro Pro Val Asn Thr His Gln Glu 20 25 30 Gln Met Lys Gln Pro Thr Pro Leu Pro Pro Pro Cys Gln Lys Val Pro 35 40 45

Page 20

Val Glu Leu Pro Val Glu Val Pro Ser Lys Gln Glu Glu Lys His Met 50 60

Thr Ala Val Lys Gly Leu Pro Glu Gln Glu Cys Glu Gln Gln Gln Lys 65 70 75 80 Glu Pro Gln Glu Gln Glu Leu Gln Gln Gln His Trp Glu Gln His Glu 85 90 95 Glu Tyr Gln Lys Ala Glu Asn Pro Glu Gln Gln Leu Lys Gln Glu Lys 100 105 110 Thr Gln Arg Asp Gln Gln Leu Asn Lys Gln Leu Glu Glu Glu Lys Lys 115 120 125 Leu Leu Asp Gln Gln Leu Asp Gln Glu Leu Val Lys Arg Asp Glu Gln 130 140 Leu Gly Met Lys Lys Glu Gln Leu Leu Glu Leu Pro Glu Gln Gln Glu 145 150 160 Gly His Leu Lys His Leu Glu Gln Gln Glu Gly Gln Leu Lys His Pro Glu Gln Gln Glu Gly Gln Leu Glu Leu Pro Glu Gln Gln Gly Gln Leu Glu Leu Pro Glu Gln Gln Glu Gly Gln Leu Glu Leu Pro Glu Gln Gln Glu Gly Gln Leu Glu Leu Pro Glu Gln Gln Glu Gly Gln Leu Glu 210 215 220 Leu Pro Gln Gln Glu Gly Gln Leu Glu Leu Ser Glu Gln Gln Glu Gly Gln Leu Glu Leu Ser Glu Gln Gln Glu Gly Gln Leu Glu Leu Ser Glu Gln Gln Glu Gly Gln Leu Lys His Leu Glu His Gln Glu Gly Gln Leu Glu Val Pro Glu Glu Gln Met Gly Gln Leu Lys Tyr Leu Glu Gln 275 280 285 Gln Glu Gly Gln Leu Lys His Leu Asp Gln Gln Glu Lys Gln Pro Glu Leu Pro Glu Gln Gln Met Gly Gln Leu Lys His Leu Glu Gln Gln Glu 315

Gly Gln Pro Lys His Leu Glu Gln Gln Glu Gly Gln Leu Glu Gln Leu Glu Glu Gln Glu Gly Gln Leu Lys His Leu Glu Gln Gln Glu Gly Gln 340 345 350 Leu Glu His Leu Glu His Gln Glu Gly Gln Leu Gly Leu Pro Glu Gln Gln Val Leu Gln Leu Lys Gln Leu Glu Lys Gln Gln Gln Pro Lys His Leu Glu Glu Glu Gly Gln Leu Lys His Leu Val Gln Gln Glu Gly Gln Leu Lys His Leu Val Gln Gln Glu Gly Gln Leu Glu Gln Gln Glu Arg Gln Val Glu His Leu Glu Gln Gln Val Gly Gln Leu Lys His Leu Glu Glu Gln Glu Gly Gln Leu Lys His Leu Glu Gln Gln Gly Gln Leu Glu Val Pro Glu Gln Gln Val Gly Gln Pro Lys Asn Leu Glu 450 Gln Glu Glu Lys Gln Leu Glu Leu Pro Glu Gln Glu Gly Gln Val 465 Lys His Leu Glu Lys Gln Glu Ala Gln Leu Glu Leu Pro Glu Gln Gln 485 490 495 Val Gly Gln Pro Lys His Leu Glu Gln Gln Glu Lys His Leu Glu His Pro Glu Gln Gln Asp Gly Gln Leu Lys His Leu Glu Gln Gln Glu Gly Gln Leu Lys Asp Leu Glu Gln Gln Lys Gly Gln Leu Glu Gln Pro Val Phe Ala Pro Ala Pro Gly Gln Val Gln Asp Ile Gln Pro Ala Leu Pro 545 550 560 Thr Lys Gly Glu Val Leu Leu Pro Val Glu His Gln Gln Gln Lys Gln
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 Asp Pro Asn Ile Gln Ala Val Arg Thr Gln Glu Lys Glu Gln Ile Lys
85 90 95
 Thr Leu Asn Asn Lys Phe Ala Ser Phe Ile Asp Lys Val Arg Phe Leu
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Ser Arg Ser Thr Ser Phe Arg Gly Gly Met Gly Ser Gly Gly Leu Ala 50 60

Thr Gly Ile Ala Gly Gly Leu Ala Gly Met Gly Gly Ile Gln Asn Glu 65 70 75 80

Lys Glu Thr Met Gln Ser Leu Asn Asp Arg Leu Ala Ser Tyr Leu Asp $85 \hspace{1.5cm} 90 \hspace{1.5cm} 95$

Arg Val Arg Ser Leu Glu Thr Glu Asn Arg Arg Leu Glu Ser Lys Ile 100 105 110

Arg Glu His Leu Glu Lys Lys Gly Pro Gln Val Arg Asp Trp Ser His 115 120 125

Tyr Phe Lys Ile Ile Glu Asp Leu Arg Ala Gln Ile Phe Ala Asn Thr 130 135

Val Asp Asn Ala Arg Ile Val Leu Gln Ile Asp Asn Ala Arg Leu Ala Page 25 Ala Asp Asp Phe Arg Val Lys Tyr Glu Thr Glu Leu Ala Met Arg Gln
165 170 175 Ser Val Glu Asn Asp Ile His Gly Leu Arg Lys Val Ile Asp Asp Thr 180 185 190 Asn Ile Thr Arg Leu Gln Leu Glu Thr Glu Ile Glu Ala Leu Lys Glu 195 200 205 Glu Leu Leu Phe Met Lys Lys Asn His Glu Glu Glu Val Lys Gly Leu 210 215 220 Gln Ala Gln Ile Ala Ser Ser Gly Leu Thr Val Glu Val Asp Ala Pro 225 230 235 240 Lys Ser Gln Asp Leu Ala Lys Ile Met Ala Asp Ile Arg Ala Gln Tyr 245 250 255 Asp Glu Leu Ala Arg Lys Asn Arg Glu Glu Leu Asp Lys Tyr Trp Ser 260 265 270 Gln Gln Ile Glu Glu Ser Thr Thr Val Val Thr Thr Gln Ser Ala Glu 275 280 285 Val Gly Ala Ala Glu Thr Thr Leu Thr Glu Leu Arg Arg Thr Val Gln 290 295 300 Ser Leu Glu Ile Asp Leu Asp Ser Met Arg Asn Leu Lys Ala Ser Leu 305 310 320 Glu Asn Ser Leu Arg Glu Val Glu Ala Arg Tyr Ala Leu Gln Met Glu 325 330 335 Gln Leu Asn Gly Ile Leu Leu His Leu Glu Ser Glu Leu Ala Gln Thr 340 345 350 Arg Ala Glu Gly Gln Arg Gln Ala Gln Glu Tyr Glu Ala Leu Leu Asn 355 360 365 Ile Lys Val Lys Leu Glu Ala Glu Ile Ala Thr Tyr Arg Arg Leu Leu 370 380 Glu Asp Gly Glu Asp Phe Asn Leu Gly Asp Ala Leu Asp Ser Ser Asn 385 395 400

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Gly Cys Phe Gly Gly Ser Ser Gly Gly Tyr Gly Gly Leu Gly Gly Phe 65 70 75 80

Gly Gly Gly Ser Phe Arg Gly Ser Tyr Gly Ser Ser Ser Phe Gly Gly $85 \hspace{1cm} 90 \hspace{1cm} 95$

Ser Tyr Gly Gly Ser Phe Gly Gly Gly Ser Phe 100 Ser Phe 105 110

Gly Gly Gly Ser Phe Gly Gly Gly Gly Phe Gly Gly Gly Phe Gly 115 120

Gly Gly Phe Gly Gly Phe Gly Gly Asp Gly Gly Leu Leu Ser Gly 130 135 140

Asn Glu Lys Val Thr Met Gln Asn Leu Asn Asp Arg Leu Ala Ser Tyr 145 150 155 160

Leu Asp Lys Val Arg Ala Leu Glu Glu Ser Asn Tyr Glu Leu Glu Gly 165 170 175

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Gly Ser Ser Gly Gly Gly Tyr Gly Gly Gly His Gly Gly Ser Ser Gly
485 490 495
Gly Gly Tyr Gly Gly Gly Ser Ser Gly Gly Gly Ser Ser Gly Gly 500 505 510
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Gly Gly His Gly Gly Ser Ser Ser Gly Gly Tyr Gly Gly Gly Ser Ser 530 540
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Met Gly Tyr Asp Asp Leu Asp Tyr Gly Met Met Ser Asp Tyr Gly Thr 290 295 300 Ala Arg Arg Thr Gly Thr Pro Ser Asp Pro Arg Arg Leu Arg Ser 305 315 320 Tyr Glu Asp Met Ile Gly Glu Glu Val Pro Ser Asp Gln Tyr Tyr Trp 325 330 335 Ala Pro Leu Ala Gln His Glu Arg Gly Ser Leu Ala Ser Leu Asp Ser 340 350 Leu Arg Lys Gly Gly Pro Pro Pro Pro Asn Trp Arg Gln Pro Glu Leu 355 360 365 Pro Glu Val Ile Ala Met Leu Gly Phe Arg Leu Asp Ala Val Lys Ser 370 380 Asn Ala Ala Ala Tyr Leu Gln His Leu Cys Tyr Arg Asn Asp Lys Val 385 390 395 400 Lys Thr Asp Val Arg Lys Leu Lys Gly Ile Pro Val Leu Val Gly Leu 405 410 415 Leu Asp His Pro Lys Lys Glu Val His Leu Gly Ala Cys Gly Ala Leu 420 425 430 Lys Asn Ile Ser Phe Gly Arg Asp Gln Asp Asn Lys Ile Ala Ile Lys 435 440 445Asn Cys Asp Gly Val Pro Ala Leu Val Arg Leu Leu Arg Lys Ala Arg 450 455 460 Asp Met Asp Leu Thr Glu Val Ile Thr Gly Thr Leu Trp Asn Leu Ser 465 470 475 480 Ser His Asp Ser Ile Lys Met Glu Ile Val Asp His Ala Leu His Ala 485 490 495 Leu Thr Asp Glu Val Ile Ile Pro His Ser Gly Trp Glu Arg Glu Pro 500 505 510 Asn Glu Asp Cys Lys Pro Arg His Ile Glu Trp Glu Ser Val Leu Thr 515 520 525 Asn Thr Ala Gly Cys Leu Arg Asn Val Ser Ser Glu Arg Ser Glu Ala 530 540 Page 33

Arg Arg Lys Leu Arg Glu Cys Asp Gly Leu Val Asp Ala Leu Ile Phe 545 550 555 560 Ile Val Gln Ala Glu Ile Gly Gln Lys Asp Ser Asp Ser Lys Leu Val 565 570 575 Glu Asn Cys Val Cys Leu Leu Arg Asn Leu Ser Tyr Gln Val His Arg 580 585 590 Glu Ile Pro Gln Ala Glu Arg Tyr Gln Glu Ala Ala Pro Asn Val Ala 595 600 605 Asn Asn Thr Gly Pro His Ala Ala Ser Cys Phe Gly Ala Lys Lys Gly 610 620 Lys Asp Glu Trp Phe Ser Arg Gly Lys Lys Pro Ile Glu Asp Pro Ala 625 635 640 Asn Asp Thr Val Asp Phe Pro Lys Arg Thr Ser Pro Ala Arg Gly Tyr 645 650 655 Glu Leu Leu Phe Gln Pro Glu Val Val Arg Ile Tyr Ile Ser Leu Leu 660 665 670 Lys Glu Ser Lys Thr Pro Ala Ile Leu Glu Ala Ser Ala Gly Ala Ile 675 680 685 Gln Asn Leu Cys Ala Gly Arg Trp Thr Tyr Gly Arg Tyr Ile Arg Ser 690 695 700 Ala Leu Arg Gln Glu Lys Ala Leu Ser Ala Ile Ala Asp Leu Leu Thr 705 710 715 720 Asn Glu His Glu Arg Val Val Lys Ala Ala Ser Gly Ala Leu Arg Asn 725 730 735 Leu Ala Val Asp Ala Arg Asn Lys Glu Leu Ile Gly Lys His Ala Ile 740 745 750 Pro Asn Leu Val Lys Asn Leu Pro Gly Gly Gln Gln Asn Ser Ser Trp 765 760 765 Asn Phe Ser Glu Asp Thr Val Ile Ser Ile Leu Asn Thr Ile Asn Glu 770 780 Val Ile Ala Glu Asn Leu Glu Ala Ala Lys Lys Leu Arg Glu Thr Gln 785 790 795 800 Page 34

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Pro Val His Asp Ala Ala Arg Glu Gly Phe Leu Asp Thr Leu Val Val 85 90 95

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Pro Ala Ala Val Ala Leu Val Leu Met Leu Leu Arg Ser Gln Arg Leu 85 90 95

Gly Gln Gln Pro Leu Pro Arg Arg Pro Gly His Asp Asp Gly Gln Arg 100 105 110

Pro Ser Gly Gly Ala Ala Ala Pro Arg Arg Gly Ala Gln Leu Arg 115 120 125

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